Docket No.: UMY-034

(PATENT)

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Patent Application of:

Mario Stevenson et al.

Application No.: 10/722,689

Confirmation No.: 3913

Filed: November 24, 2003

Art Unit: 1635

For:

MODULATION OF HIV REPLICATION BY

RNA INTERFERENCE

Examiner: Richard A. Schnizer

MS Amendment Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT (SIDS)

Dear Sir:

In accordance with 37 CFR 1.97, Applicants hereby make of record the following additional documents. A PTO Form SB/08 and a full copy of each of the documents required under 37 CFR 1.98(a)(2) accompany this statement.

This statement is not to be interpreted as a representation that the cited documents are material, that an exhaustive search has been conducted, or that no other relevant information exists. Nor shall the citation of any document herein be construed *per se* as a representation that such document is prior art. Moreover, Applicants understand the Examiner will make an independent evaluation of the cited documents.

Application No.: 10/722,689 Docket No.: UMY-034

In accordance with 37 CFR §1.97(c)(2) and §1.17(p), please charge the \$180.00 submission fee to our Deposit Order Account No. 12-0080. Please charge any necessary additional fees or credit any overpayments to our Deposit Order Account No. 12-0080.

Dated: July 16, 2007

DJM/JHV/jcb

Respectfully submitted,

Debra J. Milasincic, Esq.

Registration No.: 46,931

James H. Velema

Registration No.: 56,130

LAHIVE & COCKFIELD, LLP

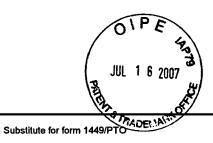
One Post Office Square

Boston, Massachusetts 02109-2127

(617) 227-7400

(617) 742-4214 (Fax)

Attorney/Agent For Applicant



INFORMATION DISCLOSURE STATEMENT BY APPLICANT

(Use as many sheets as necessary)

| Sheet | 1 | of | |
|-------|---|----|--|

| Complete if Known | | | | |
|------------------------|------------------------|--|--|--|
| Application Number | 10/722,689-Conf. #3913 | | | |
| Filing Date | November 24, 2003 | | | |
| First Named Inventor | Mario STEVENSON | | | |
| Art Unit | 1635 | | | |
| Examiner Name | Richard A. Schnizer | | | |
| Attorney Docket Number | UMY-034 | | | |

| | | | U.S. PA | TENT DOCUMENTS | | |
|-----------------------|--------------------------|--|--------------------------------|--|---|--|
| Examiner Initials* | Cite No. ¹ | Document Number Number-Kind Code ² (if known) | Publication Date MM-DD-YYYY | Name of Patentee or Applicant of Cited Document | Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear | |
| | A1* | US-6,426,073 | 07-30-2002 | Alizon et al. | | |
| | A2* | US-20020086356-A1 | 07-04-2002 | Tuschl et al. | | |
| | A3* | US-20030055020-A1 | 03-20-2003 | Fire et al. | | |
| | A4* | US-20030068821 | 04-10-2003 | Lois-Caballe et al. | | |
| | A5* | US-20030101472-A1 | 05-29-2003 | Baltimore et al. | | |
| | A6* | US-20030124513-A1 | 07-03-2003 | McSwiggen | | |
| | A7* | US-20030175950-A1 | 09-18-2003 | McSwiggen | | |
| | A8* | US-20040192629-A1 | 09-30-2004 | Xu et al. | | |
| | A9* | US-20040229266-A1 | 11-18-2004 | Tuschl et al. | | |
| | A10* | US-20040259248-A1 | 12-23-2004 | Tuschl et al. | | |
| | A11* | US-20050026278-A1 | 02-03-2005 | Tuschl et al. | | |
| | A12* | US-20050181382-A1 | 08-18-2005 | Zamore et al. | | |
| | A13* | US-20050191618-A1 | 09-01-2005 | McSwiggen et al. | | |
| | A14* | US-20050234007-A1 | 10-20-2005 | Tuschl et al. | | |
| | A15* | US-20070003960-A1 | 01-04-2007 | Tuschl et al. | | |
| | A16* | US-20070003963-A1 | 01-04-2007 | Tuschl et al. | | |
| | A17* | US-7,195,916-A1 | 03-27-2007 | Qin et al. | | |
| | | US-20030051263-A1 | 03-13-2003 | Fire et al. | | |
| | A19* | US-20030056235-A1 | 03-20-2003 | Fire et al. | | |
| | A20* | US-20030101471-A1 | 05-29-2003 | Baltimore et al. | | |
| | A21* | US-20030108923-A1 | 06-12-2003 | Tuschl et al. | | |
| | A22* | US-20030157691-A1 | 08-21-2003 | Qin et al. | | |
| | A23* | US-20040006035-A1 | 01-08-2004 | Macejak et al. | | |
| | A24* | US-20040203145-A1 | 10-14-2004 | Zamore et al. | | |
| | A25* | US-20040259247-A1 | 12-23-2004 | Tuschl et al. | | |
| | A26* | US-20050020525-A1 | 01-27-2005 | McSwiggen et al. | | |
| | A27* | US-20050037988-A1 | 02-17-2005 | Zamore et al. | | |
| | A28* | US-20050186586-A1 | 08-25-2005 | Zamore et al. | | |
| | A29* | US-20050234006-A1 | 10-20-2005 | Tuschl et al. | | |
| | | US-20060009402-A1 | 01-12-2006 | Zamore et al. | | |
| | A31* | US-20070003962-A1 | 01-04-2007 | Tuschl et al. | | |

| | FOREIGN PATENT DOCUMENTS | | | | | | | | |
|-----------------------|--------------------------|--|-----------------------------------|--|---|----|--|--|--|
| Examiner Initials* | Cite No. ¹ | Foreign Patent Document Country Code ³ -Number ⁴ -Kind Code ⁵ (if known) | Publication Date MM-DD-YYYY | Name of Patentee or Applicant of Cited Document | Pages, Columns, Lines, Where Relevant Passages Or Relevant Figures Appear | T⁵ | | | |
| | B1 | WO-02/44321-A2 | 06-06-2002 | Maxplanck-Gesellschaft | | | | | |
| | B2 | WO-2004/014933-A1 | 02-19-2004 | University of Massachusetts | | | | | |
| | B3 | WO-2004/111191-A2 | 12-23-2004 | University of Massachusetts | | | | | |
| | B4 | WO-2004/042027-A2 | 05-21-2004 | University of Massachusetts | | П | | | |

| Examiner | Date |
|-----------|------------|
| Signature | Considered |
| | |

| Sub | stitute for form 1449/PTO | | | Complete if Known | | |
|-------|---------------------------|----------|------------|------------------------|------------------------|--|
| | | | | Application Number | 10/722,689-Conf. #3913 | |
| l IN | IFORMATIO | N DI | SCLOSURE | Filing Date | November 24, 2003 | |
| S | STATEMENT BY APPLICANT | | | First Named Inventor | Mario STEVENSON | |
| | | | | Art Unit | 1635 | |
| } | (Use as many si | heets as | necessary) | Examiner Name | Richard A. Schnizer | |
| Sheet | 2 | of | 4 | Attorney Docket Number | UMY-034 | |

| B5 | WO-01/75164-A2 | 10-11-2001 | Whitehead Institute for Biomedical Research | |
|--------|-------------------|------------|---|---------------|
| B6 | WO-2005/001043-A2 | 01-06-2005 | University of Massachusetts | $\overline{}$ |
| B7 | WO-03/006477-A1 | 01-23-2003 | University of Massachusetts | |
| B8 | WO-03/022052-A1 | 03-20-2003 | California Institute of Technology | |
| В9 | WO-2003/070193-A2 | 08-28-2003 | Sirna Therapeutics, Inc. | |
| B10 | WO-03/022228-A2 | 03-20-2003 | California Institute of Technology | |
| B11 | WO-03/102131-A2 | 12-11-2003 | Sirna Therapeutics, Inc. | |
| B12 | WO-03/022040-A2 | 03-20-2003 | California Institute of Technology | |
| B13 | WO-02/097114-A2 | 12-05-2002 | Ribozyme Pharmaceuticals, Inc. | |
| B14 | WO-2005/019453-A2 | 03-03-2005 | Sirna Therapeutics, Inc. | |

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. *CITE NO.: Those application(s) which are marked with an single asterisk (*) next to the Cite No. are not supplied (under 37 CFR 1.98(a)(2)(iii)) because that application was filed after June 30, 2003 or is available in the IFW. ¹ Applicant's unique citation designation number (optional). ² See Kinds Codes of USPTO Patent Documents at www.uspto.gov or MPEP 901.04. ³ Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). ⁴ For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. ⁵ Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST.16 if possible. ⁶ Applicant is to place a check mark here if English language Translation is attached.

| | | NON PATENT LITERATURE DOCUMENTS | |
|----------------------|--------------------------|---|----|
| Examiner Initials | Cite No. ¹ | Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published. | T² |
| | C1 | Arteaga, Jose et al., "Choosing CCR5 or Rev siRNA in HIV-1," Nature, Vol. 21:230-31 (2003) | |
| | C2 | Bernstein, Emily et al., "Role for a bidentate ribonuclease in the initiation step of RNA interference," <i>Nature</i> , Vol. 409:363-366 (2001) | |
| | C3 | Billy, Eric et al., "Specific interference with gene expression induced by long, double-stranded RNA in mouse embryonal teratocarcinoma cell lines," PNAS, Vol. 98(25):14428-14433 (2001) | |
| | C4 | Bitko, Vira et al., "An Endoplasmic Reticulum-Specific Stress-Activated Caspase (Caspase-12) is Implicated in the Apoptosis of A549 Epithelial Cells by Respiratory Syncytial Virus," <i>Journal of Cellular Biology</i> , Vol. 80: 441-454 (2001) | |
| | C5 | Bitko, Vira et al., "Phenotypic silencing of cytoplasmic genes using sequence-specific double-stranded short interfering RNA and its application in the reverse genetics of wild type negative-strand RNA," <i>BMC Microbiology</i> , Vol. 1(34): 1-11 (2001) | |
| | C6 | Bowerman, Bruce et al., "A nucleoprotein complex mediates the integration of retroviral DNA," Genes & Development, Vol. 3:469-478 (1989) | |
| | C7 | Brichacek, Beda et al., "Quantitative Competitive RNA PCR for Quantitation of Virion-Associated HIV-1 RNA," <i>Methods</i> , Vol. 12:294-299 (1997) | |
| | C8 | Caplen, Natasha et al., "dsRNA-mediated gene silencing in cultured Drosophila cells: a tissue culture model for the analysis of RNA interference," <i>Gene</i> , Vol. 252:95-105 (2000) | |
| | C9 | Caplen, Natasha et al., "Specific inhibition of gene expression by small double stranded RNAs | |

| Examiner | Date | |
|-----------|------------|--|
| Signature | Considered | |
| | | |

| Sub | Substitute for form 1449/PTO | | | Complete if Known | | |
|------------------------|-----------------------------------|----|----------------------|------------------------|-------------------|--|
| | | | Application Number | 10/722,689-Conf. #3913 | | |
| INFORMATION DISCLOSURE | | | SCLOSURE | Filing Date | November 24, 2003 | |
| S | STATEMENT BY APPLICANT | | First Named Inventor | Mario STEVENSON | | |
| | | | | Art Unit | 1635 | |
| Ì | (Use as many sheets as necessary) | | Examiner Name | Richard A. Schnizer | | |
| Sheet | 3 | of | 4 | Attorney Docket Number | UMY-034 | |

| | in invertebrate and vertebrate systems," PNAS, Vol. 98(17):9742-9747 (2001) | | | | |
|------|---|--|--|--|--|
| C10 | Coburn, Glen et al., "Potent and Specific Inhibition of Human Immunodeficiency Virus Type 1 Replication by RNA Interference," <i>Journal of Virology</i> , Vol. 76(18):9225-9231 (2002) | | | | |
| C11 | Dornburg, Ralph et al., "HIV-1 Gene Therapy: Promise for the Future," <i>Advances in Pharmacology</i> , Vol. 49:229-261 (2000) | | | | |
| C12 | | | | | |
| C13. | Elbashir, Sayda et al., "Functional anatomy of siRNAs for mediating efficient RNAi in Drosophila melanogaster embryo lysate," <i>The EMBO Journal</i> , Vol. 20(23):6877-6888 (2001) | | | | |
| C14 | Elbashir, Sayda et al., "RNA interference is mediated by 21- and 22-nucleotide RNAs," Genes & Development, Vol. 15:188-200 (2001) | | | | |
| C15 | Fire, Andrew et al., "Potent and specific genetic interference by double-stranded RNA in Caenorhabditis elegans," Nature, Vol. 391:806-811 (1998) | | | | |
| C16 | Hammond, Scott et al., "An RNA-directed nuclease mediates post-transciptional gene silencing in Drosophila cells," <i>Nature</i> , Vol. 404:293-296 (2000) | | | | |
| C17 | Hutvágner, György et al., "RNAi:nature abhors a double-strand," Current Opinion in Genetics & Development, Vol. 12:225-232 (2002) | | | | |
| C18 | Ketting, René et al., "mut-7 of C. elegans, Required for Transposon Silencing and RNA Interference, Is a Homolog of Werner Syndrome Helicase and RNaseD," <i>Cell</i> , Vol. 99:133-141 (1999) | | | | |
| C19 | Kimpton, Jaculyn et al., "Detection of Replication-Competent and Pseudotyped Human Immunodeficiency Virus with a Sensitive Cell Line on the Basis of Activation of an Integrated ß-Galactosidase Gene," <i>Journal of Virology</i> , Vol. 66(4):2232-2239 (1992) | | | | |
| C20 | Li, Yuexia et al., "Molecular Characterization of Human Immunodeficiency Virus Type 1 Cloned Directly from Uncultured Human Brain Tissue: Identification of Replication-Competent and -Defective Viral Genomes," <i>Journal of Virology</i> , Vol. 65(8):3973-3985 (1991) | | | | |
| C21 | Moore, John et al., "New Targets for Inhibitors of HIV-1 Replication," <i>Nature</i> , Vol. 1:40-49 (2000) | | | | |
| C22 | Nykänen, Antti et al., "ATP Requirements and Small Interfering RNA Structure in the RNA Interference Pathway," Cell, Vol. 107:309-321 (2001) | | | | |
| C23 | Paddison, Patrick et al., "Stable suppression of gene expression by RNAi in mammalian cells," PNAS, Vol. 99(3):1443-1448 (2002) | | | | |
| C24 | Sharkey, Mark et al., "Persistence of episomal HIV-1 infection intermediates in patients on highly active anti-retroviral therapy," <i>Nature Medicine</i> , Vol. 6(1):76-81 (2000) | | | | |
| C25 | Sharp, J.G. et al., "Oligonucleotide Enhanced Cytotoxicity of Idarubicin for Lymphoma Cells," Leukemia and Lymphoma, Vol. 42:417-427 (2001) | | | | |
| C26 | Sharp, Philip. "RNA interference-2001," Genes and Development, Vol. 15:485-490 (2001) | | | | |
| C27 | Tabara, Hiroaki et al., "The <i>rde-1</i> Gene, RNA Interference and Transpoon Silencing in <i>C. elegans</i> ," <i>Cell</i> , Vol. 99:123-132 (1999) | | | | |
| C28 | Ui-Tei, Kumiko et al., "Sensitive assay of RNA interference in <i>Drosophila</i> and Chinese hamster cultured cells using firefly luciferase gene as target," <i>FEBS Letters</i> , Vol. 479:79-82 (2000) | | | | |
| C29 | Wang, Ming-Bo et al., "High-efficiency silencing of a ß-glucuronidase gene in rice is correlated with repetitive transgene structure but is independent of DNA methylation," <i>Plant Molecular Biology</i> , Vol. 43:67-82 (2000) | | | | |

| Examiner | Date | |
|-----------|------------|---|
| Signature | Considered | · |

| Substitute for form 1449/PTO | | | | Complete if Known | | | |
|-----------------------------------|-----------|------|-----------|------------------------|------------------------|--|--|
| | | | | Application Number | 10/722,689-Conf. #3913 | | |
| 11 | NFORMATIC | N DI | SCLOSURE | Filing Date | November 24, 2003 | | |
| S | TATEMENT | BY A | APPLICANT | First Named Inventor | Mario STEVENSON | | |
| (Use as many sheets as necessary) | | | | Art Unit | 1635 | | |
| | | | | Examiner Name | Richard A. Schnizer | | |
| Sheet | 4 | of | 4 | Attorney Docket Number | UMY-034 | | |

| C30 | Welker, Reinhold et al., "Virion Incorporation of Human Immunodeficiency Cirus Type 1 Nef Is Mediated by a Bipartite Membrane-Targeting Signal: Analysis of Its Role in Enhancement of Viral Infectivity," <i>Journal of Virology</i> , Vol. 72(11):8833-8840 (1998) | |
|-----|--|--|
| C31 | Wesley, S. Varsha et al., "Construct design for efficient, effective and high-throughput gene silencing in plants," <i>The Plant Journal</i> , Vol. 27(6):581-590 (2001) | |
| C32 | Yang, Shicheng et al., "Specific Double-Stranded RNA Interference in Undifferentiated Mouse Embryonic Stem Cells," <i>Molecular and Cellular Biology</i> , Vol. 21(22):7807-7816 (2001) | |
| C33 | Zamore, Phillip et al., "RNAi: Double-Stranded RNA Directs the ATP-Dependent Cleavage of mRNA at 21 to 23 Nucleotide Intervals," Cell, Vol. 101:25-33 (2000) | |
| C34 | International Search Report for Application No. PCT/US03/37860 dated 7/21/2005 | |
| C35 | Written Opinion for Application No. PCT/US03/37860 dated 5/31/2007 | |

^{*}EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

| Examiner | D | ate | |
|-----------|---|-----------|--|
| Signature | | onsidered | |
| | | | |

¹Applicant's unique citation designation number (optional). 2Applicant is to place a check mark here if English language Translation is attached.